

JAP20 Rec'd PSTIPTO 20 JUL 2008

SEQUENCE LISTING

<1	.10>	AKZ	O No	bel	N.V.											
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Glł	a ctt / Leu	tta Leu 15	a agt ı Sei	cat His	Gly	gaa Glu	a aaq 1 Lys 20	g ato s Met	g aat : Asr	gtt Val	ati	ggt Gl _y 25	aat Asr	aad Asr	c ata n Ile	99
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cat His 125	gga Gly	tat Tyr	gct Ala	ctt Leu	cag Gln 130	ggt Gly	tgg Trp	aaa Lys	att Ile	gat Asp 135	aat Asn	act Thr	gaa Glu	G1 y 999	cca Pro 140	435
caa	cgt	atc	tca	ggt	ggt	gtt	aat	cça	ggt	aca	aat	act	tcg	cag	att	483

Glr	a Arg	; Ile	e Sei	Gly 145		/ Val	. Asr	Pro	Gly 150		Ası	ı Thi	: Sei	Gl: 15	ı Ile	
at <u>o</u> Met	ggt Gly	aca Thr	ggt Gly 160	/ Glu	cca Pro	aca Thr	gat Asp	ato Ile 165	Arg	ctt Lei	gat 1 Asp	act Thr	tgg Trp	Thi	gtt Val	531
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act Thr	ggt Gly	gtc Val	Pro	ggt Gly 385	aaa Lys	aat Asn	aca Thr	Ala	gga Gly 390	tct Ser	gtt Val	ggt Gly	gat Asp	gct Ala 395	acc Thr	1203
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tct ggg Ser Gly	ttt Phe	ggt Gly 640	tca Ser	att a Ile 1	aac Asn .	Ala	aat Asn 645	act Thr	tta Leu	gaa Glu	Gly	tca Ser 650	aac Asn	gta Val	1971
gat ata Asp Ile	tct Ser 655	aca Thr	gag (Glu)	ttt q Phe N	Val :	tca Ser: 660	atg (Met	att (gca Ala	Thr	caa Gln 665	cgt Arg	gga Gly	ttc Phe	2019

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Thr Thr Asn Val Ser Phe Asn Val Asn Leu Ser Ser Asp Lys Ser Gly

185

180

- Asp Lys Ser Gln Asn Val Asn Ser Pro Phe Thr Ser Leu Phe Asn Ile 195 200 205
- Trp Asn Gly Lys Gln Pro Ser Glu Pro Asn Asn Pro Pro Met Pro Glu 210 215 220
- Ser Ala Tyr Ser Tyr Gln Thr Ser Ile Lys Val Tyr Asp Glu Ala Gly 225 230 235 240
- Gly Thr His Thr Leu Thr Val Tyr Phe Asp Gln Val Ser Pro Lys Asp 245 250 255
- Tyr Lys Gly Gly Gly Ser Gly Glu Ser Val Trp Glu Tyr Val Val Thr 260 265 270
- Met Asp Pro Ser Glu Asp Asn Arg Gln Val Ser Val Gly Gly Asn Ile 275 280 285
- Val Asp Ile Lys Asp Thr Lys Ala Ala Gly Met Leu Met Ser Gly Thr 290 295 300
- Leu Ser Phe Asp Ser Ser Gly Lys Leu Ala Asn Gln Ser Ala Tyr Ser 305 310 315 320
- Leu Asn Gly Ser Arg Lys Pro Ala Val Asp Pro Ala Thr Gly Ala Leu 325 330 335
- Ile Asn Gly Asn Gly Phe Thr Ile Asp Arg Asp Gly Asn Ala Ile Pro
 340 345 350
- Ile Leu Asn Ile Asp Asn Pro Ala Glu Asn Phe Tyr Pro Ala Glu Val 355 360 365
- Ser Asn Asn Gly Phe Pro Met Ile Val Ala Asn Phe Thr Gly Val Pro 370 380
- Gly Lys Asn Thr Ala Gly Ser Val Gly Asp Ala Thr Thr Phe Phe Thr 385 390 395 400
- Glu Ile Asp Phe Gly Leu Lys Ala Thr Asp Leu Asp Asn Thr Trp Lys 405 410 415
- Asn Ala Asn Glu Pro Leu Ser Ser Leu Ser Tyr Lys Lys Thr His Asn 420 425 430
- Pro Met Asp Val Ala Gly Gly Trp Thr Val Gly Gly Tyr Lys Thr Pro 435 440 445
- Ala Pro Ser Val Thr Glu Leu Gly Met Ala Gln Ile Leu Glu Asn Pro

450 455 460

Ala Gly Val Met Pro Gln Tyr Tyr Phe Gly Asn Pro Asn Tyr Asp Asn 465 470 475 480

Thr Val Pro Gln Ser Pro Pro Tyr Val Tyr Lys Asn Glu Ala Ser Tyr 485 490 495

Gln Ala Ala Tyr Lys Thr Ala Leu Thr Ala Ala Gly Gly Thr Ala Ala 500 505 510

Asp Ile Lys Lys Glu His Trp Pro His Asn Ala Ala Ser Gly Ile Leu 515 520 525

Glu Ala Asn Asp Pro Pro Asn Val Lys Asp Leu Ala Asn Met Asn Gly 530 540

Thr Pro Asn Arg Leu Ser Asn Ala Phe Thr Asn Tyr Ala Gly Gly Ser 545 550 560

Ser Thr Lys Ser Ala Ser Gln Asn Gly Tyr Gly Phe Gly Asp Leu Met 565 570 575

Asn Tyr Ser Val Asn Ala Glu Gly Val Leu Phe Gly Val Tyr Ser Asn 580 585 590

Gly Val Gln Leu Pro Leu Tyr Gln Val Ala Leu Tyr Asp Phe Asn Ser 595 600 605

Lys Gln Gly Leu Arg Arg Glu Gly Gly Asn Leu Phe Ser Gln Thr Arg 610 615 620

Glu Ser Gly Asp Pro Ser Ser Gly Ala Ala Asn Thr Ser Gly Phe Gly 625 630 635 640

Ser Ile Asn Ala Asn Thr Leu Glu Gly Ser Asn Val Asp Ile Ser Thr 645 650 655

Glu Phe Val Ser Met Ile Ala Thr Gln Arg Gly Phe Gln Ser Asn Ser 660 665 670

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gag ttc ata caa gat ttt ccc aat agc tat caa gaa gat gga caa atg Glu Phe Ile Gln Asp Phe Pro Asn Ser Tyr Gln Glu Asp Gly Gln Met 25 30 35	148
gtt aca gga att att tca aaa ata ata ggc tct aac tgt gat aat tct Val Thr Gly Ile Ile Ser Lys Ile Ile Gly Ser Asn Cys Asp Asn Ser 40 45 50 55	196
tca aca tct gat ata aat aat aag aaa tcc ata gat aga gat aaa gat Ser Thr Ser Asp Ile Asn Asn Lys Lys Ser Ile Asp Arg Asp Lys Asp 60 65 70	244
aca tta ctc tca agt agt aat aga aat aca ata caa gcc ggt act cca Thr Leu Leu Ser Ser Ser Asn Arg Asn Thr Ile Gln Ala Gly Thr Pro 75 80 85	292
cat caa gaa aat aac ata aaa gaa gat ctt caa ctg act aac aaa aat His Gln Glu Asn Asn Ile Lys Glu Asp Leu Gln Leu Thr Asn Lys Asn 90 95 100	340
gaa caa aca act cca gaa gaa gaa gaa agt aaa ttt att tgg tta Glu Gln Thr Thr Pro Glu Glu Glu Glu Glu Ser Lys Phe Ile Trp Leu 105 110 115	388
aca gaa gct cca tca gag ctt aaa aaa gga gaa aaa gct ata aca caa Thr Glu Ala Pro Ser Glu Leu Lys Lys Gly Glu Lys Ala Ile Thr Gln 120 125 130 135	436
aca aga ttg tct att ggt aag gat ata tct ttt aga att act gct gat Thr Arg Leu Ser Ile Gly Lys Asp Ile Ser Phe Arg Ile Thr Ala Asp 140 145 150	484
gat gcc atc aaa gct caa tca atg atg tta aaa aat cca gat agg ttt Asp Ala Ile Lys Ala Gln Ser Met Met Leu Lys Asn Pro Asp Arg Phe 155 160 165	532
gtt tta gat ctt caa gga aag tgg ggt att tcc ctt cca cct att cca Val Leu Asp Leu Gln Gly Lys Trp Gly Ile Ser Leu Pro Pro Ile Pro 170 175 180	580
CCT aca aat CCT tgg tta aaa aaa ata Cgc tta ggt act aat aat gga Pro Thr Asn Pro Trp Leu Lys Lys Ile Arg Leu Gly Thr Asn Asn Gly 185 190 195	628
aat aca cga ctt gtc ttt gat ctt caa aaa aaa cca tct aaa act gaa Asn Thr Arg Leu Val Phe Asp Leu Gln Lys Lys Pro Ser Lys Thr Glu 200 205 210 215	676

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Tyr Gln Glu Asp Gly Gln Met Val Thr Gly Ile Ile Ser Lys Ile Ile 35 40 45	
Gly Ser Asn Cys Asp Asn Ser Ser Thr Ser Asp Ile Asn Asn Lys Lys 50 55 60	
Ser Ile Asp Arg Asp Lys Asp Thr Leu Leu Ser Ser Ser Asn Arg Asn 65 70 75 80	
Thr Ile Gln Ala Gly Thr Pro His Gln Glu Asn Asn Ile Lys Glu Asp 85 90 95	
Leu Gln Leu Thr Asn Lys Asn Glu Gln Thr Thr Pro Glu Glu Glu Glu 100 105 110	
Glu Ser Lys Phe Ile Trp Leu Thr Glu Ala Pro Ser Glu Leu Lys Lys 115 120 125	
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Ser Phe Arg Ile Thr Ala Asp Asp Ala Ile Lys Ala Gln Ser Met Met 145 150 155 160	
Leu Lys Asn Pro Asp Arg Phe Val Leu Asp Leu Gln Gly Lys Trp Gly 165 170 175	
Ile Ser Leu Pro Pro Ile Pro Pro Thr Asn Pro Trp Leu Lys Lys Ile 180 185 190	
Arg Leu Gly Thr Asn Asn Gly Asn Thr Arg Leu Val Phe Asp Leu Gln 195 200 205	

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WO 2005/070958 PCT/EP2005/000562 9

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agc tat act tat ttg gaa aaa gct atc cca tct atg tta act tct aga Ser Tyr Thr Tyr Leu Glu Lys Ala Ile Pro Ser Met Leu Thr Ser Arg 40 45 50 55	198
ctt tat tgg gaa gaa cgt ttt caa cct atc ccg gat gct aat gct att Leu Tyr Trp Glu Glu Arg Phe Gln Pro Ile Pro Asp Ala Asn Ala Ile 60 65 70	246
aaa gca gga aag gta gaa gat ata aag gaa atg gat aag gca agg ata Lys Ala Gly Lys Val Glu Asp Ile Lys Glu Met Asp Lys Ala Arg Ile 75 80 85	294
gct aca ggt gca gac tat ctt ata tgg gga cag gta aat att gta ggt Ala Thr Gly Ala Asp Tyr Leu Ile Trp Gly Gln Val Asn Ile Val Gly 90 95 100	342
gat gaa gct acg ctt gat gta caa gtt tgt gat ata gaa gga tca att Asp Glu Ala Thr Leu Asp Val Gln Val Cys Asp Ile Glu Gly Ser Ile 105 110 115	390
tgg agg a.aa agt aaa aat aca aaa gtt gat aat tta att act gcc ctt Trp Arg Lys Ser Lys Asn Thr Lys Val Asp Asn Leu Ile Thr Ala Leu 120 125 130 135	438
caa gat aca gca gat gca att aat agt gag ttg ttt ggg cgt gca act Gln Asp Thr Ala Asp Ala Ile Asn Ser Glu Leu Phe Gly Arg Ala Thr 140 145 150	486
aca aaa cca tca tca aaa gct act att gta gct caa atg aac tct gga Thr Lys Pro Ser Ser Lys Ala Thr Ile Val Ala Gln Met Asn Ser Gly 155 160 165	534
ttg att aæg gga aaa gga aat gaa aat cag tca tat ctt aat cca gaa	582

Leu Ile Lys Gly Lys Gly Asn Glu Asn Gln Ser Tyr Leu Asn Pro Glu ttt cgt tat caa gga agc aat ctt tcc cgt ggc cga agt caa gct ctt 630 Phe Arg Tyr Gln Gly Ser Asn Leu Ser Arg Gly Arg Ser Gln Ala Leu 190 ccc ttt gct tca gtt ggt ata gtt gtt ggt gac ttt ata gga gat aat 678 Pro Phe Ala Ser Val Gly Ile Val Val Gly Asp Phe Ile Gly Asp Asn 205 aaa aat gaa gtt gcc ata tta agt gag tat aaa gtc cat att tat cga 726 Lys Asn Glu Val Ala Ile Leu Ser Glu Tyr Lys Val His Ile Tyr Arg 774 tgg gaa gaa agg tta gct ctt ctt gga gaa tat aaa ttc cct cgc Trp Glu Glu Glu Arg Leu Ala Leu Leu Gly Glu Tyr Lys Phe Pro Arg 240 tca cta cag tct tta cat att cgt gct ttt gat gtg gat cat gat ggt 822 Ser Leu Gln Ser Leu His Ile Arg Ala Phe Asp Val Asp His Asp Gly gta cag gaa atc att gtt tct tgc ttt gat cct tca tat gca aag cca 870 Val Gln Glu Ile Ile Val Ser Cys Phe Asp Pro Ser Tyr Ala Lys Pro 270 265 tat tcg ttt att ctt agt ttt aaa aat aga gtg ttt aaa gag tta gcc 918 Tyr Ser Phe Ile Leu Ser Phe Lys Asn Arg Val Phe Lys Glu Leu Ala 285 aca aac tta cca ttt tat tta aat gtg gtt aaa ctt cca cca gat ttt 966 Thr Asn Leu Pro Phe Tyr Leu Asn Val Val Lys Leu Pro Pro Asp Phe 305 300 tot cot atg tta att ggt caa aag agt gac aat tca agg att ttt tct Ser Pro Met Leu Ile Gly Gln Lys Ser Asp Asn Ser Arg Ile Phe Ser 315 ccc tct ggg gtt tat gaa ata gaa aaa cat gga cgt aac tat ata atg 1062 Pro Ser Gly Val Tyr Glu Ile Glu Lys His Gly Arg Asn Tyr Ile Met 335 gga aat cgt ctt agt ctt cca aag gaa gct aat att ttt aat ttt tct Gly Asn Arg Leu Ser Leu Pro Lys Glu Ala Asn Ile Phe Asn Phe Ser 350 tgg tta cca tca gat tca tta aaa gat gaa gaa gct aag tta gta ctt 1158 Trp Leu Pro Ser Asp Ser Leu Lys Asp Glu Glu Ala Lys Leu Val Leu 365 gta acc aat aat gaa aga tta gtt gta tat aat aca aaa ggt aca aga 1206 Val Thr Asn Asn Glu Arg Leu Val Val Tyr Asn Thr Lys Gly Thr Arg 380 ctt ttt atg act gaa gaa gtg tat tat ggt tct tct gtt ggt ata gac Leu Phe Met Thr Glu Glu Val Tyr Tyr Gly Ser Ser Val Gly Ile Asp 395 gag ccc agt aat atg cct ggt ctt gga aag tca aaa gag ctt atc cct 1302 Glu Pro Ser Asn Met Pro Gly Leu Gly Lys Ser Lys Glu Leu Ile Pro tct aaa tat ttt atc cca gga cgg atg att cct att aat ctt gat tca 1350 Ser Lys Tyr Phe Ile Pro Gly Arg Met Ile Pro Ile Asn Leu Asp Ser

425		430		435		
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tta aca tgg (Leu Thr Trp i			y Leu Val		_	_
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gga cta gaæ. 2 Gly Leu Glu I 520			-			_
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12 .

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Ala Asn Ile Phe Asn Phe Ser Trp Leu Pro Ser Asp Ser Leu Lys Asp

13

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Gly Ser Ser Val Gly Ile Asp Glu Pro Ser Asn Met Pro Gly Leu Gly 410

Lys Ser Lys Glu Leu Ile Pro Ser Lys Tyr Phe Ile Pro Gly Arg Met 425

Ile Pro Ile Asn Leu Asp Ser Met Gly Lys Trp Glu Leu Leu Val Ser 440

Lys Pro Ile Ser Val Ala Ala Lys Phe Phe Glu Asn Tyr Arg Ser Phe 450 455

Ala Glu Gly Glu Ile Gln Ala Leu Thr Trp Asp Gly Leu Gly Leu Gly 470

Leu Val Trp Asn Thr Arg Arg Ile Lys Gly Thr Ile Thr Asp Phe Ala 490

Leu Ala Asp Met Asn Asn Asp Gly Lys Leu Asp Leu Val Val Ser Val 505

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			actt Leu													19:	9
			tta Lev													24	7
			tto Phe													299	5
			tgg Trp												Pro	343	3
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			act Th <u>r</u>													439	,
tta Leu	gaa Glu 135	gaa Glu	atc Ile	gct Ala	cgt Arg	tca Ser 140	aca Thr	aaa Lys	act Thr	gta Val	gtc Val 145	att Ile	gct Ala	cta Leu	agt Ser	487	•
			aat Asn													535	
			gaa Glu													583	
			cgg Arg 185													631	
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			gct Ala													775	
			gca Ala					Lys								823	
aca	tat	gga	aat	aat	tgg	cct	ata	gca	ttt	tat	cca	tat	tat	aga	ttt	871	

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		Asp										aag Lys				967
	Ile											gat Asp				1015
	-			_				_	_			gga Gly			_	1063
	_		_									tta Leu				1111
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Glu Lys Glu Pro Leu Gln Tyr Ile Leu Leu Lys Lys Glu Lys Phe Ser 50 55 60

Gln Met Glu Ile Tyr Asn Tyr Gln Phe Thr Ser Gln His Trp Ser Pro 65 70 75 80

Asp Asn Phe Val Ser Pro Ala Ile Trp Glu His Gln Val Asp Ile Tyr 85 90 95

Ile Pro His His Pro Val Ser Glu Arg Ala Leu Leu Ile Ile Asn Asn 100 105 110

Gly Ile Asn Asn Gly Thr Phe Phe Thr Ser Pro Lys Ala Pro Thr Asp 115 120 125

Phe Thr Pro Glu Val Leu Glu Glu Ile Ala Arg Ser Thr Lys Thr Val

Val Ile Ala Leu Ser Asp Ile Pro Asn Gln Tyr Leu Thr Tyr Arg Gly 145 150 155 160

Asp Trp Arg Phe Leu Lys Glu Asp Glu Ser Ile Ala Met Ser Trp Ser 165 170 175

Ser Phe Leu Gln Asp Pro Glu Ser Arg Tyr Thr Arg Pro Leu Tyr Val 180 185 190

Pro Met Val Ala Æla Val Ser Gln Ala Met Thr Leu Ala Glu Lys Glu 195 200 205

Leu Gln Ala Leu Lys Ile Lys His Phe Ile Val Ser Gly Væl Ser Lys 210 215 220

Arg Gly Trp Thr Thr Trp Leu Ser Ala Ile Ala Asp Ser Arg Val Asp 225 230 235 240

Ala Ile Thr pro Phe Val Ile Asp Ala Leu Asn Thr Arg Ly's Val Leu

245 250 255

Gly His Met Tyr Lys Thr Tyr Gly Asn Asn Trp Pro Ile Ala Phe Tyr 260 265 270

Pro Tyr Tyr Arg Phe Asp Leu Asp Lys Gln Leu Asp Thr Val Pro Phe 275 280 285

Phe Asn Leu Met Asn Ile Val Asp Pro Tyr Arg Tyr Leu Gly Thr Pro 290 295 300

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Asp Asp Phe Tyr Val Pro Asp Asn Ser Ser Phe Tyr Tyr Asp Asp Leu 325 330 335

Pro Gly Glu Lys Ala Leu Arg Phe Ala Pro Asn Ser Asn His His Gly 340 345 350

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Gln Lys Gly Ile Ser Thr Pro Val Leu Asp Ile Ser Thr Glu Met Thr 370 375 380

Glu Arg Val Gln Tyr Val Thr Val Arg Phe Ser Glu Val Pro Glu Lys 385 390 395 400

Ile Val Leu Trp Lys Ala Ala Asn Pro Glu Ser Arg Asp Phe Arg Tyr 405 410 415

Ala Cys Arg Val Arg Tyr Met Glu Thr Pro Leu His Leu Ser Ala Thr 420 425 430

Gly Glu Val Ser Val Ser Leu Glu Ile Pro Ser Val Gly Trp Gln Ala 435 440 445

Ala Phe Ile Glu Ala Thr Phe Lys Asp Gly Phe Val Ala Thr Thr Pro 450 455 460

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aca Thr	aaa Lys	aaa Lys	tat Tyr	aaa Lys 50	gca Ala	ttt Phe	gaa Glu	gag Glu	ctt Leu 55	ata Ile	aaa Lys	aca Thr	gtt Val	aaa Lys 60	gat Asp	194
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aca Thr 110	aac Asn	acg Thr	att Ile	ctt Leu	tct Ser 115	aat Asn	aat Asn	cat His	att Ile	ttt Phe 120	gat Asp	tct Ser	aaa Lys	act Thr	gaa Glu 125	386
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	Arg							ggc Gly								866
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							Gly	att Ile 470								1442
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Ser Leu Ser Lys Leu Ser Ser Val Gly Ala Ile Leu Lys Lys Glu Gly 65 70 75 80

Ser Val Ser Asn Thr Ser Val Ala Ser Val Lys Ala Ser Ser Asp Ala 85 90 95

Ser Asp Gly Thr His Thr Ile Asp Val Lys Gln Leu Ala Thr Asn Thr 100 105 110

Ile Leu Ser Asn Asn His Ile Phe Asp Ser Lys Thr Glu Ser Ile Asn 115 120 125

Asn Thr Gly Ser Pro Gly Ile Phe Ala Tyr Glu Tyr Lys Gly Glu Leu 130 135 L40

His Glu Val Glu Val Pro Pro Gly Ser Asp Leu G-lu Tyr Leu Ala Thr 145 150 155 160

Leu Ile Asn Lys Asp Ser Asn Asn Pro Gly Val Lys Ala Asn Leu Ile 165 170 175

Lys Thr Gly Asp Gly Tyr Met Phe Ser Leu Glu Gly Thr Glu Thr Gly 180 185 190

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Gly Arg Thr Phe Thr Phe Asp Ile Pro Ser Gly Thir Thr Ala Lys Glu 245 250 255

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Lys Lys Gln Val Glu Lys Ala Val Glu Ser Ile Asn Asn Val Leu Ser 370 375 380

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Lys Gln Cys Thr Thr Ala Leu Ser Arg Trp Lys Glu Lys His Lys Gly

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Gly Gly Glu Asn Ser Thr Leu Thr Pro Leu Glu Met Met Arg Glu Leu

280

295

275

290

867

915

300

PCT/EP2005/000562

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Ile Asp Asn Ala Ser Glu Glu Leu Thr Thr Ser Leu Glu Ser Lys Se 50 55 60	r
	r
Ser Asp Asp Phe Ala Ile Lys Asp Arg Lys Arg Gln Gly Lys Gly Se 65 70 75 80	
Asp Ser Leu Leu Lys Met Val Gln Glu Tyr Thr Glu Leu Thr Asn As	p

- Asn Lys Thr Asp Ala Trp Ala Ser Leu Ala Glu Ile Ala Glu Tyr 130 135 140
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- Met Glu Thr Leu Glu Asn Glu Phe Gly Asp Glu Ile Lys Ala Gly Leu 165 170 175
- Lys Gly Ala Leu Asn Ser Lys Glu Phe Thr Asp Ile Gly Ser Ala Ala 180 185 190
- Gln Leu Arg Asp Leu Tyr Thr Thr Thr Val Thr Ile Thr Ala Ala Pro 195 200 205
- Asp Ala Val Leu Ala Arg Leu Leu Glu Glu Tyr Glu Ser Asp Asp Asp 210 215 220
- Leu Asp Arg Ala Ile Asp Phe Leu Leu Ser Thr Leu Gly Gly Glu Leu 225 230 235 240
- Glu Ser Ala Asp Pro Ser Met Asp Lys Val His Leu Gln Ser Val Met 245 250 255
- Gly Asp Ile Glu Lys Thr Gln Gln Leu His Ser Ser His Lys Gln Cys 260 265 270
- Thr Thr Ala Leu Ser Arg Trp Lys Glu Lys His Lys Gly Gly Glu Glu 275 280 285
- Asn Ser Thr Leu Thr Pro Leu Glu Met Met Arg Glu Leu Il e Ala Leu 290 295 300
- Lys Asn Glu Asn Phe Ile Ser Pro Ser Ser Ile Asp Lys Ile Val Asp 305 310 315 320
- Gln Ala Asp Pro Gln Asp Ile Glu Lys Glu Val Leu Phe Leu Gln Glu
 325 330 335
- Met Leu Ala Ala Val Arg Lys Phe Pro Ile Met Val Phe Asp Asn Val 340 345 350
- Glu Asn Arg Val Arg Val Met Gly Ala Val Gln Asp Ala Val Asp Asp 355 360 365
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					gat Asp 45											196
					tca Ser											244
					gta Val											292
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					att Ile											532
					ctt Leu											580
					gct Ala											628

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									tcg Ser 225	Leu						724
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gca Ala	gaa Glu	aca Thr 250	gca Ala	caa Gln	tgg Trp	att Ile	gca Ala 255	ttt Phe	ggrt Gly	ata Ile	cag Gln	tta Leu 260	gca Ala	atg Met	att Ile	820
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gct Ala 280	gga Gly	gtg Val	tca Ser	aaa Lys	ata Ile 285	gca Ala	gat Asp	atg Met	tt t Ph e	tca Ser 290	aag Lys	tct Ser	caa Gln	gat Asp	gta Val 295	916
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gct Ala	ggt Gly	tca Ser	gtt Val 315	aat Asn	cag Gln	tct Ser	gct Ala	ata Ile 320	gg⊂ Gly	ggt Gly	aca Thr	ggt Gly	att Ile 325	ggt Gly	aca Thr	1012
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									gct. Ala							1108
									ttc Phe							1156
ata Ile	aca Thr	gat Asp	Ile	att Ile 380	caa Gln	gat Asp	agc Ser	gtc Val	aat Asn 385	aca Thr	aaa Lys	ata Ile	Ala	gtt Val 390	caa Gln	1204
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- Leu Met Thr Gly Thr Ser Gly Glu Glu Glu Glu Glu Glu Glu Gly 25 25 30
- Gly Val Pro Lys Glu Gln Gly Gly Pro Gly Lys Gly Asp Ala Ser Glu 35 40 45
- Ala Ala Lys Gly Gln Ala Ala Ala Asp Ser Ile Asn Ser Ala Gly Gly 50 55 60
- Thr Glu Lys Pro Gly Glu Val Gly Asp Lys Glu Asp Val Gly Glu Gly 65 70 75 80
- Gly Glu Ile Pro Glu Gly Gly Glu Ile Pro Glu Gly Gly Glu Glu Val 85 90 95
- Pro Glu Glu Pro Pro Tyr Val Pro Pro Pro Leu Val Glu Pro Ala Lys 100 \$105\$
- Ile Ser Thr Val Thr Asp Leu Ser Thr Leu Met Gly Ser Leu Gln Leu 115 120 125
- Thr Glu Gln Lys Lys Asn Ala Glu Lys Thæ Val Asn Glu Ile Lys Ala 130 135 140
- Glu Asp Asn Ile Ala Glu Ser Lys Lys Ser Gly Ile Leu Lys Phe Phe 165 170 175
- Gln Lys Leu Phe Ala Val Ile Gly Ala Val Leu Gly Ala Ile Gly Gly 180 185 190
- Ala Leu Ala Ile Ala Ala Gly Ala Ala Ser Gly Asn Pro Leu Leu Val 195 200 205
- Ala Ala Gly Ile Met Ala Ile Val Ala Ser Ile Asp Ala Ala Met Ser 210 215 220
- Ser Leu Ser Asp Gly Lys Val Ser Ile Ser Ala Gly Ile Ser Lys Ala 225 230 235 240
- Leu Glu Ala Met Gly Val Pro Ala Glu Thr Ala Gln Trp Ile Ala Phe 245 250 255
- Gly Ile Gln Leu Ala Met Ile Ala Val Thr Ile Ala Ile Gly Phe Ala 260 265 270

Ser Gly Gly Gly Ala Met Ala Gly Val Ser Lys Ile Ala Asp Met Phe Ser Lys Ser Gln Asp Val Ala Lys Leu Ala Gln Met Ile Glu Lys Ala Ser Lys Ile Val Gln Ile Ala Gly Ser Val Asn Gln Ser Ala Ile 315 Gly Gly Thr Gly Ile Gly Thr Ala Val Val Gln Ser Asn Ile Lys Ala 330 Asn Glu Ser Glu Gln Lys Glu Ile Glu Ala Ala Ile Ala Lys Val Lys Ala Lys Ile Glu Thr Leu Gln Asp Phe Phe Lys Asn Gln Met Glu Gln Phe Asn Ala Ile Met Lys Ile Ile Thr Asp Ile Ile Gln Asp Ser Val 375 Asn Thr Lys Ile Ala Val Gln Arg Gly Ala Arg Glu <210> 15 <211> 894 <212> DNA <213> Lawsonia intracellularis <220> <221> CDS <222> (13)..(894) aggaggaatt at atg tot ott gto att aat aac aac otg atg goo gto aat 51 Met Ser Leu Val Ile Asn Asn Leu Met Ala Val Asn get caa egt aac tta age aag tet tat gga gaa etg agt tet tet gtt 99 Ala Gln Arg Asn Leu Ser Lys Ser Tyr Gly Glu Leu Ser Ser Ser Val 20 cga aaa ctt tct tca ggt ctt cgt gta gga act gct gct gat gac tca 147 Arg Lys Leu Ser Ser Gly Leu Arg Val Gly Thr Ala Ala Asp Asp Ser 35 gca ggg tta gcc att cga gaa ctc atg aga tct gac att gca aca aca 195 Ala Gly Leu Ala Ile Arg Glu Leu Met Arg Ser Asp Ile Ala Thr Thr 50 55 caa caa gga ata cga aat gcg aat gct att tct atq att caa act 243 Gln Gln Gly Ile Arg Asn Ala Asn Asp Ala Ile Ser Met Ile Gln Thr 65 gcg gat ggt gca ctt gga gtc atc gat gaa aag ctc att cga atg aaa 291

Ala	Asp	Gly 80	Ala	Leu	Gly	Val		Asp	Glu	Lys	Leu		Arg	Met	Lys	
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	att					ta t Tyr	caa				Ser			act Thr		387
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cgt Arg	atc Ile	cac His 160	ttt Phe	G1 y 999	aca Thr	tct Sex	aac Asn 165	agc Ser	tct Ser	gct Ala	gaa Glu	gat Asp 170	tac Tyr	tat Tyr	gat Asp	531
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														gat Asp		723
tta Leu	Ser	ata Ile 240	caa Gln	aat Asn	gaa Glu	aac Asn	tta Leu 245	caa Gln	gct Ala	gct Ala	gaa Glu	tct Ser 250	cgt Arg	ata Ile	tct Ser	771
Asp						gaa Glu 260								caa Gln	ata Ile	819
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Ser Ser Gly Leu Arg Val Gly Thr Ala Ala Asp Asp Ser Ala Gly Leu 35 40 45

Ala Ile Arg Glu Leu Met Arg Ser Asp Ile Ala Thr Thr Gln Gln Gly 50 60

Ile Arg Asn Ala Asn Asp Ala Ile Ser Met Ile Gln Thr Ala Asp Gly 65 70 75 80

Ala Leu Gly Val Ile Asp Glu Lys Leu Ile Arg Met Lys Glu Leu Ala 85 90 95

Glu Gln Ala Ala Thr Gly Thr Tyr Asn Ser Thr Gln Arg Met Ile Ile 100 105 110

Asp Ser Glu Tyr Gln Ala Met Ala Ser Glu Ile Thr Arg Ile Ala Asn 115 120 125

Ala Thr Glu Phe Asn Gly Ile Lys Leu Leu Asp Gly Ser Leu Ser Gly
130 135 140

Asn His Asp Gly Lys Lys Ile Asn Ser Thr Gly Ala Val Arg Ile His 145 150 155 160

Phe Gly Thr Ser Asn Ser Ser Ala Glu Asp Tyr Tyr Asp Ile Lys Ile 165 170 175

Gly Gly Ser Thr Ala Ser Ala Leu Gly Leu Gly Asn Thr Val Lys Gly
180 185 190

Ala Gly Ala Thr Val Ser Thr Gln Ala Ala Ala Gln Asn Ala Leu Lys
195 200 205

Ala Ile Asp Asn Ala Ile Val Ser Lys Asp Lys Ile Arg Ala His Leu 210 215 220

Gly Gly Leu Gln Asn Arg Leu Glu Ala Thr Val Asp Asn Leu Ser Ile 225 230 235 240

Gln Asn Glu Asn Leu Gln Ala Ala Glu Ser Arg Ile Ser Asp Ile Asp
245 250 255

Val Ser Gln Glu Met Thr Gln Phe Val Arg Asn Gln Ile Leu Thr Gln 260 265 270

Thr Gly	/ Val / 275	Ala Me	t Leu	Ser	Gln 280	Ala	Asn	Sex	Leu	Pro 285	Arg	Met	Ala
Gln Glr 29 C		Ile Gl	Y										

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Asn	Val 170	Gln	Thr	Pro	Leu	Leu 175	Ile	Gly	Ser	Gly	Val 180	Ser	Asn	Gly	Pro	
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				_				gga Gly	_	_	_	_				676
		-				_		aaa Lys 225			_					724
		_						aat Asn					_		-	772
								gta Val								820
								aat Asn								868
								ctt Leu								916
		_	_					ata Ile 305		_			-	_		964
								gta Val		_			_		_	1012
								gaa Glu								1060
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					Gly			cgt Arg								1300
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425					430					435					440	
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cca Pro	gaa Glu 490	ctt Leu	cca Pro	tat Tyr	tta Leu	gtt Val 495	aac Asn	ata Ile	caa Gln	aaa Lys	ggt Gly 500	gct Ala	atg Met	aca Thr	caa Gln	1540
tgg Trp 505	ggc Gly	act Thr	aaa Lys	aat Asn	atg Met 510	ccc Pro	ttt Phe	tta Leu	ttg Leu	gat Asp 515	aca Thr	aga Arg	aat Asn	tta Leu	atc Ile 520	1588
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tta Leu	ata Ile ·	cct Pro 555	gly ggg	aac Asn	caa Gln	cct Pro	gtt Val 560	ata Ile	cag Gln	ttt Phe	ctt Leu	gga Gly 565	ggt Gly	cct Pro	caa Gln	1732
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cca Pro 585	aaa Lys	ctt Leu	att Ile	gga Gly	agt Ser 590	agc Ser	atg Met	gtg Val	cca Pro	ctt Leu 595	gct Ala	ttt Phe	gtc Val	tct Ser	caa Gln 600	1828
tct Ser	ttt Phe	tca Ser	tca Ser	cca Pro 605	gat Asp	ctt Leu	ttt Phe	gtt Val	aaa Lys 610	caa Gln	act Thr	aga Arg	agt Ser	ggt Gly 615	ctc Leu	1876 ·
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						aga Arg 655										2020
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gtt Val	caa Gln	ggt Gly	gga Gly	cat His 685	gaa Glu	agt Ser	ctt Leu	gat Asp	act Thr 690	tct Ser	att Ile	ggt Gly	agc Ser	aaa Lys 695	gca Ala	2116

				atg Met											2164
			_	atg Met		-			-		_			_	2212
	Gly		_	aaa Lys			-							_	2260
 a Glu			_	gtt Val 750	_			-							2308
			_	aca Thr	_							-			2356
				Gly 333			_						_		2404
				ctt Leu											2452
		-		cct Pro	-						_				2500
His			_	tta Leu 830					_	_	_			_	2548
_	_			aca Thr	_						_	_		_	2596
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	_			aac Asn		_		_			_			_	2692
				aat Asn											2740
Asn				gga Gly 910											2788
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aaa Lys		taa													2848

37

WO 2005/070958 PCT/EP2005/000562

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Asn Lys Pro Phe Asp Ser Phe Phe Gly Gly Ser Asp Ser Thr Ile Gly

Thr Leu Glu Thr Gly Pro Thr Asn Leu Thr Phe Thr Thr Val Gly Ala

Phe Arg Asn Ser Val Phe Arg Ile Ile Gly Gly Gly Arg Ser Ser Phe

Asn Asn Pro Asn Thr Val Lys Gly Asn Val Thr Leu Thr Val Tyr Asn 105

Thr Asp Val Glu Arg Ile Ile Gly Ala Gly Ile Ser Asn Arg Gly Leu

Val Thr Val Thr Gly Ser Val Asn Met Lys Leu Glu Asn Val Ser Val

Thr Arg Gly Ile Tyr Gly Gly Val Tyr Thr Gln Asn Gly His Val Leu

Gly Ser Ile Asn Met His Leu Lys Asn Val Gln Thr Pro Leu Leu Ile 170

Gly Ser Gly Val Ser Asn Gly Pro Asn Arg Ile Thr Val Asn Gly Asp 185

Ile Asn Ile Asp Val Glu Asp Ser Arg Ile Gln Tyr Val Asn Ile Thr

Gly Glu Val Asp Ala Gly Ile Lys Gly Asn Ala Thr Leu Thr Val Lys

Lys Ser Thr Val Glu Leu Ile Asn Ser Gly Arg Gly Asn Ile Leu Gly

225 230 235 240 Asn Leu Lys Ile Ser Ile Ala Asp Ser Asn Ile Arg Gly Leu Ser Pro 250 Val Asp Phe Gly Ser Ser Val Tyr Gly Asp Thr Ser Ile Asn Val Ile 260 265 Asn Ser Gln Ile Asn Asp Ile Thr Leu Ile Pro Arg Ala Gly Gly Met 280 Leu Val Gly Pro Val Thr Leu Asp Ile Thr Ser Ser Thr Ile Gln Asn 295 Ile Gln Cys Gly Pro Val Ser Gln Asn Asn Gln Leu Asn Thr Leu Asn 315 Val Thr Val Asn Thr Ser Asn Ile Thr Asn Leu Asn Leu Gly Ser Val 330 Glu Gly His Thr Ile Ser Thr Thr Ala Thr Val Thr Asp Ser Asn Ile 340 345 Thr Asn Leu Asn Val Gly Thr Phe Asn Gly Leu Gly Val Thr Glu Asn 360 Ala Ser Val Ile Ile Asn Ser Gly Asn Ile Thr Asn Leu Asn Val Gly 375 Thr Asn Val Ile Ala Ala Ala Thr Thr Ile Asn Ser Ser Ala Thr Ile 390 395 His Asp Gly Leu Ile Ala Asn Leu Thr Leu Gly Ser Gln Gly Asn Gly 405 Arg Thr Met Ile Ala Thr Ala Asn Val Asn Gly Gly Thr Ile Gly Leu 425

Glu Leu Ala Ile Leu Asn Met Ser Gly Gly Leu Ile Glu Arg Ile Ile 455

Leu Thr Met Gly Ser Glu Asn Phe Ile Pro Gly Thr Arg Pro Ile Thr 440

Val Gly Asn Ala Asn Ser Ser Thr Ile Asn Phe Thr Pro Gly Lys Arg 470 475

Ser Ile Val Lys Thr Ile Asn Gly Pro Glu Leu Pro Tyr Leu Val Asn 485 490

Ile Gln Lys Gly Ala Met Thr Gln Trp Gly Thr Lys Asn Met Pro Phe

505

500

Leu Leu Asp Thr Arg Asn Leu Ile Leu Ser Gly Thr Leu Ile Thr Ser 515 520 525

Asn Ile Gln Leu Ala Asp Leu Ser Ile Thr Asn Leu Phe Val Ala Asn 530 540

Gly Gly Thr Leu Val Pro Arg Lys Leu Ile Pro Gly Asn Gln Pro Val 545 550 560

Ile Gln Phe Leu Gly Gly Pro Gln Ser Leu Leu Val Ile His Gln Pro 565 570 575

Leu Lys Val Asn Leu Ser Leu Ser Pro Lys Leu Ile Gly Ser Ser Met 580 585 590

Val Pro Leu Ala Phe Val Ser Gln Ser Phe Ser Ser Pro Asp Leu Phe 595 600 605

Val Lys Gln Thr Arg Ser Gly Leu Ile Trp Ser Asp Leu Glu Phe Asp 610 615 620

Pro Thr Thr Ser Ile Trp Tyr Val Asn Asn Ile Gln Ala Ser Gln Asp 625 630 635 640

Phe Tyr Ser Phe Ser Ile Ala Arg Glu Thr Thr Asn Trp Leu Arg Gln 645 650 655

Gln His Ile Trp Thr Leu Gln Asn Arg Ser Ser Lys Leu Leu Asp Asn 660 665 670

Glu His Tyr Gly Leu Trp Ile Asn Val Gln Gly Gly His Glu Ser Leu 675 680 685

Asp Thr Ser Ile Gly Ser Lys Ala Lys Met Pro Trp Ile Met Ala Thr 690 695 700

Ala Gly Tyr Asp Tyr Leu Gln Gln Leu Pro Arg Leu Asp Met Lys Ala 705 710 715 720

Leu Tyr Gly Leu Ala Phe Gly Ala Ser Lys Gly Lys Ser Lys Trp Ser

Ser Val Asn Ser Thr Lys Asn Asp Ala Glu Leu Gly Met Val Ser Gly 740 745 750

Tyr Val Gly Leu Ile His Asn Lys Thr Gly Leu Tyr Ser Thr Leu Thr 755 760 765

Leu Gln Leu Ala Ser Ser Lys Leu His Thr Asn Ser Thr Gly Phe Tyr 770 775 780

Arg Asn Phe Lys Trp Thr Glu Thr Thr Pro Thr Glu Ala Leu Glu Leu 785 790 795 800

Gly Trp Lys Tyr Thr Phe Asn Asn Gly Ile Lys Met Asn Pro Arg Gly 805 810 815

Gln Leu Ile Phe Glu Gln Thr Ser Lys His His Phe Asp Leu Gly Ile 820 825 830

Gln Asn Asp Lys Ala Ile Leu Asp Lys Ser Gln Leu Ile Thr Ser Ser 835 840 845

Leu Gly Ile Thr Val Glu Tyr Lys Leu Pro Val Thr Thr Pro Ile Asn 850 855 860

Leu Tyr Ala Gly Ile Glu Arg Ile Lys Gly Gln Ser Gly Asn Phe Ala 865 870 875 880

Ile Ser Ser Gln Ser Leu Gln Met Lys Phe Lys His Asp Asn Asp Thr 885 890 895

Ser Val Val Arg Ala Thr Ile Gly Thr Asn Ile Leu Leu Gly Glu His 900 905 910

Phe Asn Ile His Cys Asp Ile Phe Gly Asp Lys Gly Asn Asp Lys Gly 915 920 925

Ile Gly Gly Gln Ala Gly Phe Thr Tyr Lys Phe 930 935